

WHAT IS CLAIMED IS:

- 1                   1. An isolated binding agent that competes with a monoclonal  
2 antibody MAb 292-2-3 for specific binding to human cytochrome P450 allelic variant  
3 2C9\*2 without specifically binding to human cytochrome 2C9\*1 and 2C9\*3, and that  
4 specifically inhibits 2C-catalyzed metabolism of phenanthrene by at least 50%.
- 1                   2. The binding agent of claim 1 that lacks specific binding to each of  
2 human cytochromes P450 1A1, 1A2, 2A6, 2B6, 2C8, 2C18, 2C19, 2D6, 2E1, 3A4, and  
3 3A5.
- 1                   3. The binding agent of claim 1 that specifically inhibits the enzyme  
2 activity of human cytochrome P450 allelic variant 2C9\*2 by at least 90%.
- 1                   4. The binding agent of claim 1 that is MAb 292-2-3 or a binding  
2 fragment thereof.
- 1                   5. The binding agent of claim 1 that is a monoclonal antibody.
- 1                   6. The monoclonal antibody of claim 5 that is a Fab fragment.
- 1                   7. The monoclonal antibody of claim 5 that is a mouse antibody.
- 1                   8. A cell line producing the monoclonal antibody of claim 5.
- 1                   9. The cell line of claim 8 that is a eucaryotic cell line.
- 1                   10. The cell line of claim 9 that is a procaryotic cell line.
- 1                   11. The monoclonal antibody of claim 5 comprising a light chain  
2 variable domain having at least 80% sequence identity with the light chain variable  
3 domain of a monoclonal antibody MAb 292-2-3 and a heavy chain variable domain  
4 having at least 80% sequence identity with the heavy chain variable domain of the  
5 monoclonal antibody MAb 292-2-3.
- 1                   12. The monoclonal antibody of claim 5, wherein the light chain  
2 variable domain comprises three CDR regions from the light chain of a monoclonal

3 antibody MAb 292-2-3, and the heavy chain variable domain comprises three CDR  
4 regions from the heavy chain of the monoclonal antibody MAb 292-2-3.

1 13. An isolated binding agent that competes with a monoclonal  
2 antibody MAb 763-15-5 for specific binding to the human cytochrome p450 2C9 allelic  
3 variants 2C9\*1, 2C9\*2, and 2C9\*3, and that specifically inhibits 2C-catalyzed  
4 metabolism of phenanthrene by at least 50%.

1 14. The binding agent of claim 13 that lacks specific binding to each of  
2 human cytochromes P450 1A1, 1A2, 2A6, 2B6, 2C9, 2C18, 2C19, 2D6, 2E1, 3A4, and  
3 3A5.

1 15. The binding agent of claim 13 that specifically inhibits the enzyme  
2 activity of human cytochrome P450 allelic variant 2C9\*2 by at least 90%.

1 16. The binding agent of claim 13 that is MAb 292-2-3 or a binding  
2 fragment thereof.

1 17. The binding agent of claim 13 that is a monoclonal antibody.

1 18. The monoclonal antibody of claim 17 that is a Fab fragment.

1 19. The monoclonal antibody of claim 17 that is a mouse antibody.

1 20. A cell line producing the monoclonal antibody of claim 17.

2 21. The cell line of claim 20 that is a eucaryotic cell line.

1 22. The cell line of claim 21 that is a procaryotic cell line.

1 23. The monoclonal antibody of claim 17 comprising a light chain  
2 variable domain having at least 80% sequence identity with the light chain variable  
3 domain of a monoclonal antibody MAb 763-15-5 and a heavy chain variable domain  
4 having at least 80% sequence identity with the heavy chain variable domain of the  
5 monoclonal antibody MAb 763-15-5.

1 24. The monoclonal antibody of claim 17, wherein the light chain  
2 variable domain comprises three CDR regions from the light chain of a monoclonal

3 antibody MAb 763-15-5, and the heavy chain variable domain comprises three CDR  
4 regions from the heavy chain of the monoclonal antibody MAb 763-15-5.

1 25. The binding agent of claim 13 that specifically inhibits the enzyme  
2 activity of human cytochrome P450 allelic variants 2C9\*1 and 2C9\*3 by at least 70%.

1 26. The binding agent of claim 13 that specifically inhibits the enzyme  
2 activity of human cytochrome P450 2C18 by 30%.

1 27. An isolated binding agent that competes with a monoclonal  
2 antibody MAb 763-15-20 for specific binding to the human cytochrome P450 2C9 allelic  
3 variants 2C9\*1, 2C9\*2, and 2C9\*3.

1 28. An isolated binding agent that competes with a monoclonal  
2 antibody selected from the group consisting of MAb 5-1-5 and MAb 281-1-1 for specific  
3 binding to human cytochrome P450 2C8, and that specifically inhibits 2C-catalyzed  
4 metabolism of phenanthrene by at least 50%.

1 29. The binding agent of claim 28 that lacks specific binding to each of  
2 human cytochromes P450 1A1, 1A2, 2A6, 2B6, 2C9, 2C18, 2C19, 2D6, 2E1, 3A4, and  
3 3A5.

1 30. The binding agent of claim 28 that specifically inhibits the enzyme  
2 activity of human cytochrome p450 2C8 by at least 90%.

1 31. The binding agent of claim 28 that is MAb 5-1-5 or a binding  
2 fragment thereof.

1 32. The binding agent of claim 28 that is MAb 281-1-1 or a binding  
2 fragment thereof.

1 33. The binding agent of claim 28 that is a monoclonal antibody.

1 34. The monoclonal antibody of claim 33 that is a Fab fragment.

1 35. The monoclonal antibody of claim 33 that is a mouse antibody.

1 36. A cell line producing the monoclonal antibody of claim 33.

- 1                   37.    The cell line of claim 36 that is a eucaryotic cell line.
- 1                   38.    The cell line of claim 37 that is a procaryotic cell line.
- 1                   39.    The monoclonal antibody of claim 33 comprising a light chain  
2    variable domain having at least 80% sequence identity with the light chain variable  
3    domain of a monoclonal antibody selected from the group consisting of MAb 5-1-5 and  
4    MAb 281-1-1, and a heavy chain variable domain having at least 80% sequence identity  
5    with the heavy chain variable domain of a monoclonal antibody selected from the group.
- 1                   40.    The monoclonal antibody of claim 33, wherein the light chain  
2    variable domain comprises three CDR regions from the light chain of a monoclonal  
3    antibody selected from the group, and the heavy chain variable domain comprises three  
4    CDR regions from the heavy chain of a monoclonal antibody selected from the group.
- 1                   41.    An isolated binding agent that competes with a monoclonal  
2    antibody MAb 592-2-5 for specific binding to human cytochrome P450 2C9 and 2C18,  
3    and that specifically inhibits 2C-catalyzed metabolism of phenanthrene by at least 50%.
- 1                   42.    The binding agent of claim 41 that specifically inhibits the enzyme  
2    activity of human cytochrome p450 2C9 by at least 80%.
- 1                   43.    The binding agent of claim 41 that specifically inhibits the enzyme  
2    activity of human cytochrome P450 2C18 by at least 80%.
- 1                   44.    The binding agent of claim 41 that is MAb 592-2-5 or a binding  
2    fragment thereof.
- 1                   45.    The binding agent of claim 41 that is a monoclonal antibody.
- 1                   46.    The monoclonal antibody of claim 45 that is a Fab fragment.
- 1                   47.    The monoclonal antibody of claim 45 that is a mouse antibody.
- 1                   48.    A cell line producing the monoclonal antibody of claim 45.
- 1                   49.    The cell line of claim 48 that is a eucaryotic cell line.

1                   50.    The cell line of claim 48 that is a prokaryotic cell line.

1                   51.    The monoclonal antibody of claim 45 comprising a light chain  
2    variable domain having at least 80% sequence identity with the light chain variable  
3    domain of a monoclonal antibody MAb 592-2-5 and a heavy chain variable domain  
4    having at least 80% sequence identity with the heavy chain variable domain of the  
5    monoclonal antibody MAb 592-2-5.

1                   52.    The monoclonal antibody of claim 45, wherein the light chain  
2    variable domain comprises three CDR regions from the light chain of a monoclonal  
3    antibody MAb 592-2-5 and the heavy chain variable domain comprises three CDR  
4    regions from the heavy chain of the monoclonal antibody MAb 592-2-5.

1                   53.    An isolated binding agent that competes with a monoclonal  
2    antibody MAb 5-7-5 for specific binding to a human cytochrome p450 2C family member  
3    selected from the group consisting of 2C9, 2C18, and 2C19, and that specifically inhibits  
4    2C-catalyzed metabolism of phenanthrene by at least 50%.

1                   54.    The binding agent of claim 53 that lacks specific binding to each of  
2    human cytochromes P450 1A1, 1A2, 2A6, 2B6, 2C8, 2D6, 2E1, 3A4, and 3A5.

1                   55.    The binding agent of claim 53 that specifically inhibits the enzyme  
2    activity of human cytochrome p450 2C9 by at least 90%.

1                   56.    The binding agent of claim 53 that specifically inhibits the enzyme  
2    activity of human cytochrome p450 2C18 by at least 90%.

1                   57.    The binding agent of claim 53 that specifically inhibits the enzyme  
2    activity of human cytochrome p450 2C19 by at least 90%.

1                   58.    The binding agent of claim 53 that is a monoclonal antibody.

1                   59.    The monoclonal antibody of claim 58 that is a Fab fragment.

1                   60.    The monoclonal antibody of claim 59 that is a mouse antibody.

1                   61.    The binding agent of claim 53 that is MAb 5-7-5 or a binding  
2    fragment thereof.

1                   62. A cell line producing the monoclonal antibody of claim 58.

1                   63. The cell line of claim 62 that is a eucaryotic cell line.

1                   64. The cell line of claim 62 that is a procaryotic cell line.

1                   65. The monoclonal antibody of claim 58 comprising a light chain  
2 variable domain having at least 80% sequence identity with the light chain variable  
3 domain of a monoclonal antibody MAb 5-7-5 and a heavy chain variable domain having  
4 at least 80% sequence identity with the heavy chain variable domain of the monoclonal  
5 antibody MAb 5-7-5.

1                   66. The monoclonal antibody of claim 58, wherein the light chain  
2 variable domain comprises three CDR regions from the light chain of a monoclonal  
3 antibody MAb 5-7-5 and the heavy chain variable domain comprises three CDR regions  
4 from the heavy chain of the monoclonal antibody MAb 5-7-5.

1                   67. A method of determining whether cytochrome P450 2C9\*2  
2 metabolizes a compound, comprising:  
3                   contacting the compound with cytochrome P450 2C9\*2 in the presence of  
4 varying amounts of the binding agent of claim 1; and  
5                   assaying metabolism of the compound as a function of amount of binding  
6 agent, a decrease of metabolism with amount of binding agent indicating that cytochrome  
7 P450 2C9\*2 metabolizes the compound.

1                   68. The method of claim 67, wherein the compound is contacted with  
2 cytochrome P450 2C9\*2 in a sample containing a collection of cytochrome P450  
3 enzymes including 2C9\*2.

1                   69. The method of claim 68, wherein the sample is a tissue sample.

1                   70. The method of claim 69, wherein the collection of enzymes are  
2 obtained from a cell culture expressing the enzymes.

1                   71. The method of claim 70, wherein the compound is a drug, steroid  
2 or carcinogen.

1           72.    A method of detecting cytochrome P450 2C9\*2, comprising:

2                   contacting a sample suspected of containing cytochrome P450 2C9\*2 with

3    a binding agent of claim 1; and

4                   determining whether the agent specifically binds to the sample, specific

5    binding indicating the presence of cytochrome P450 2C9\*2 in the sample.

1           73.    A method of measuring P450 2C9\*2 levels in an individual relative

2    to P450 2C9\*2 levels in a control population, the method comprising:

3                   contacting a sample suspected of containing cytochrome P450 2C9\*2 from

4    the individual with a binding agent of claim 1, and

5                   determining the P450 2C9\*2 levels in the individual relative to P450

6    2C9\*2 a mean level in a control population.